

PTO/SB/08A

**INFORMATION DISCLOSURE  
STATEMENT BY APPLICANT**

Use as many sheets as necessary)

**Complete if Known**

Application Number	09/857,873
Filing Date	October 5, 2001
Confirmation Number	Unknown
First Named Inventor	John P. McKearin et al.
Group Art Unit	1614
Examiner Name	Unknown

Sheet

1

of

12

Attorney Docket No.

PHA 4293 (3167/4Z)

**U.S. PATENT DOCUMENTS**

Examiner Initials*	Cite No. <sup>1</sup>	U.S. Patent Document		Name of Patentee or Applicant of Cited Document	Date of Publication of Cited Document MM-DD-YYYY
		Number	Kind Code <sup>2</sup> (if known)		
	1.	4,100,274		Dutta et al.	07-11-1978
	2.	4,105,681		Bollag et al.	08-08-1978
	3.	4,140,707		Cleare et al.	02-20-1979
	4.	4,215,215		Bollag et al.	07-29-1980
	5.	4,310,666		Zee-Cheng et al.	01-12-1982
	6.	4,472,382		Labrie et al.	09-18-1984
	7.	4,596,797		Schweikert et al.	06-24-1986
	8.	5,250,683		Holton et al.	10-05-1993
	9.	5,254,703		Holton	10-19-1993
	10.	5,272,171		Ueda et al.	12-21-1993
	11.	5,319,112		Kingston et al.	06-07-1994
	12.	5,344,991		Reitz et al.	09-06-1994
	13.	5,455,270		Kaplan et al.	10-03-1995
	14.	5,633,016		Johnson	05-27-1997
	15.	5,686,419		Powers et al.	11-11-1997
	16.	5,696,131		Baguley et al.	12-09-1997

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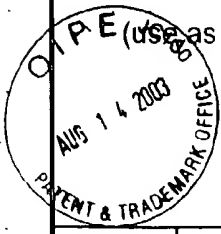
<sup>1</sup>Unique citation designation number. <sup>2</sup>See attached Kinds of U.S. Patent Documents. <sup>3</sup>Enter Office that issued the document, by the two-letter code (WIPO Standard ST.3). <sup>4</sup>For Japanese patent documents, the indication of the year of the reign of the Emperor must precede the serial number of the patent document. <sup>5</sup>Kind of document by the appropriate symbols as indicated on the document under WIPO Standard ST. 16 if possible. <sup>6</sup>Applicant is to place a check mark here if English language Translation is attached or place an "A" here if English language abstract is attached..

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PTO/SB/08A				<b>Complete if Known</b>	
<b>INFORMATION DISCLOSURE STATEMENT BY APPLICANT</b>  (use as many sheets as necessary)				Application Number	09/857,873
				Filing Date	October 5, 2000
				Confirmation Number	Unknown
				First Named Inventor	John P. McKearn et al.
				Group Art Unit	1614
				Examiner Name	Unknown
Sheet	2	of	12	Attorney Docket No.	PHA 4293 (3167/4Z)



17.	5,733,909	Black et al.	03-31-1998
18.	5,767,142	LaVoie et al.	06-16-1998
19.	5,824,699	Kreft et al.	10-20-1998
20.	5,837,696	Golub et al.	11-17-1998
21.	5,869,524	Failli	02-09-1999
22.	5,952,381	Chen et al.	09-14-1999

### FOREIGN PATENT DOCUMENTS

Examiner Initials*	Cite No. <sup>1</sup>	Foreign Patent Document			Name of Patentee or Applicant of Cited Document	Date of Publication of Cited Document MM-DD-YYYY	T <sup>6</sup>
		Office	Number <sup>4</sup>	Kind Code <sup>2</sup> (if known)			
	23.	DE	195 48 798	A1	Dr. Karl Thomae GmbH	07-03-1997	A
	24.	DE	196 13 933	A1	Merck Patent GmbH	10-09-1997	A
	25.	DE	196 26 701	A1	Hoechst AG	01-08-1998	A
	26.	EP	0 010 458	A1	ANVAR	04-30-1980	A
	27.	EP	0 054 168	A1	Klinge Pharma GmbH & Co.	06-23-1982	A
	28.	EP	0 095 875	A2	Farmos Group Ltd.	12-07-1983	
	29.	EP	0 165 904	A2	Ciba-Geigy AG	12-27-1985	A
	30.	EP	0 199 636	A1	C.I.R.D.	10-29-1986	A
	31.	EP	0 236 940	B1	Ciba-Geigy AG	09-16-1987	
	32.	EP	0 260 744	B1	Janssen Pharmaceutica N.V.	03-23-1988	

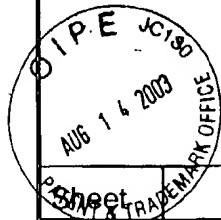
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				First Named Inventor	John P. McKearr et al.
				Group Art Unit	1614
				Examiner Name	Unknown
				Attorney Docket No.	PHA 4293 (3167/4Z)
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33.	EP	0 299 402	A2	ASTA Pharma Aktiengesellschaft	01-18-1989	A
34.	EP	0 331 983	A2	F. Hoffmann-La Roche AG	09-13-1989	A
35.	EP	0 402 232	A1	Rhone-Poulenc Sante	12-12-1990	A
36.	EP	0 532 156	A1	Matsuno et al.	03-17-1993	
37.	EP	0 579 915	A1	F. Hoffmann-La Roche AG	01-26-1994	A
38.	EP	0 702 962	A2	Eli Lilly and Company	03-27-1996	
39.	EP	0 703 239	A1	Hoechst Aktiengesellschaft	03-27-1996	A
40.	EP	0 716 086	A1	Boehringer Mannheim GMBH	06-12-1996	
41.	EP	0 743 070	A2	Trnka Mudr et al.	11-20-1996	A
42.	EP	0 758 649	A1	Kureha Chemical Industry Co., Ltd.	02-19-1997	
43.	EP	0 882 734	A2	F. Hoffman-La Roche AG	12-09-1998	
44.	EP	0 921 119	A1	ADIR Et Compagnie	06-09-1999	A
45.	GB	2 282 598	A	Merck & Co., Inc.	04-12-1995	
46.	WO	87/07609	A1	Taiho Pharmaceutical Co., Ltd.	12-17-1987	A
47.	WO	93/11145	A1	Banyu Pharmaceutical Co., Ltd.	06-10-1993	
48.	WO	93/18652	A1	Merck & Co.	09-30-1993	
49.	WO	94/12169	A1	Merck & Co., Inc.	06-09-1994	
50.	WO	94/13635	A1	Merck Frosst Canada Inc.	06-23-1994	
51.	WO	94/20480	A1	Merck Frosst Canada Inc.	09-15-1994	
52.	WO	94/21612	A1	Otsuka Pharmaceutical Co., Ltd.	09-29-1994	

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		Group Art Unit	1614	
		Examiner Name	Unknown	
Sheet	4	12	Attorney Docket No.	PHA 4293 (3167/4Z)

<input checked="" type="checkbox"/>	53.	WO	94/25431	A1	The Welcome Foundation Limited	11-10-1994	
	54.	WO	94/25466	A1	Glaxo, Inc.	11-10-1994	
	55.	WO	94/26731	A1	Merck Frosst Canada Inc.	11-24-1994	
	56.	WO	94/27980	A1	G.D. Searle & Co.	12-08-1994	
	57.	WO	95/02603	A2	Pharmacia S.P.A.	01-26-1995	
	58.	WO	95/08327	A1	The Government of the United States of America	03-30-1995	
	59.	WO	95/11883	A1	G.D. Searle & Co.	05-04-1995	
	60.	WO	95/12606	A1	The Welcome Foundation Limited	05-11-1995	
	61.	WO	95/15316	A1	G.D. Searle & Co.	06-08-1995	
	62.	WO	95/24199	A2	Cell Therapeutics, Inc.	09-14-1995	
	63.	WO	95/30652	A1	G.D. Searle & Co.	11-16-1995	
	64.	WO	95/30656	A1	G.D. Searle & Co.	11-16-1995	
	65.	WO	96/00574	A1	Smithkline Beecham Corporation	01-11-1996	
	66.	WO	96/01653	A1	Board of Regents, The University of Texas System et al.	01-25-1996	
	67.	WO	96/03387	A1	G.D. Searle & Co.	02-08-1996	
	68.	WO	96/03392	A1	G.D. Searle & Co.	02-08-1996	
	69.	WO	96/06840	A1	Merck Frosst Canada Inc.	03-07-1996	
	70.	WO	96/14745	A1	The Regents of the University of California	05-23-1996	
<input checked="" type="checkbox"/>	71.	WO	96/21667	A1	Merck Frosst Canada Inc.	07-18-1996	

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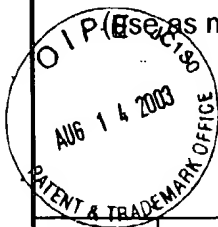
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Application Number 09/857,873  
 Filing Date October 5, 2000  
 Confirmation Number Unknown  
 First Named Inventor John P. McKearn et al.  
 Group Art Unit 1614  
 Examiner Name Unknown

Sheet 5 of 12 Attorney Docket No. PHA 4293 (3167/4Z)

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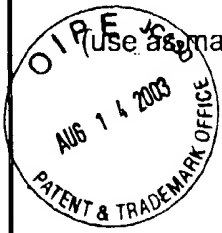
72.	WO	96/33988	A1	Wisconsin Alumni Research Foundation	10-31-1996	
73.	WO	96/35774	A2	The Children's Medical Center Corporation	11-14-1996	
74.	WO	96/36335	A1	Pharmacia S.P.A.	11-21-1996	
75.	WO	96/36612	A1	Rutgers, The State University of New Jersey	11-21-1996	
76.	WO	96/36622	A1	Indena S.P.A.	11-21-1996	
77.	WO	96/36623	A1	Merck Frosst Canada Inc.	11-21-1996	
78.	WO	96/37469	A1	Merck Frosst Canada Inc.	11-28-1996	
79.	WO	97/15666	A1	The Children's Medical Center Corporation	05-01-1997	
80.	WO	97/15676	A2	Abbott Laboratories	05-01-1997	
81.	WO	97/19954	A1	Asta Medica Aktiengesellschaft	06-05-1997	
82.	WO	97/20835	A1	Harbor Branch Oceanographic Institution, Inc.	06-12-1997	
83.	WO	97/24116	A2	Allergan	07-10-1997	
84.	WO	97/29106	A1	Rutgers, The State University of New Jersey	08-14-1997	
85.	WO	97/31936	A2	Wisconsin Alumni Research Foundation	09-04-1997	
86.	WO	97/34608	A1	Mayo Foundation for Medical Education and Research	09-25-1997	
87.	WO	97/36497	A2	Mars, Incorporated	10-09-1997	

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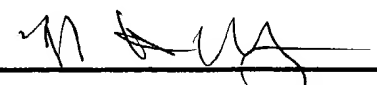
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Sheet	6	of	12	Attorney Docket No.	PHA 4293 (3167/4Z)

88.	WO	97/36863	A1	Merck Frosst Canada Inc.	10-09-1997	
89.	WO	97/37658	A1	Boehringer Mannheim Italia S.P.A.	10-16-1997	
90.	WO	97/41844	A1	Alcon Laboratories, Inc.	11-13-1997	
91.	WO	97/47296	A2	Mount Sinai Hospital Corporation	12-18-1997	
92.	WO	97/49704	A1	Janssen Pharmaceutica N.V.	12-31-1997	
93.	WO	98/03484	A1	Merck Frosst Canada Inc.	01-29-1998	
94.	WO	98/07433	A1	Bristol-Myers Squibb Company	02-26-1998	
95.	WO	98/12181	A1	Rutgers, The State University of New Jersey	03-26-1998	
96.	WO	98/13350	A1	Zeneca Limited	04-02-1998	
97.	WO	98/14188	A1	Ilex Oncology Inc.	04-09-1998	
98.	WO	98/16227	A1	G.D. Searle & Co.	04-23-1998	
99.	WO	98/22101	A2	G.D. Searle & Co.	05-28-1998	
100.	WO	98/25603	A1	Ilex Oncology, Inc.	06-18-1998	
101.	WO	98/25896	A1	G.D. Searle & Co.	06-18-1998	
102.	WO	98/40104	A2	Yale University et al.	09-17-1998	
103.	WO	98/45294	A1	Astra Pharmaceuticals Ltd.	10-15-1998	
104.	WO	99/01131	A1	Smithkline Beecham Corporation	01-14-1999	
105.	WO	99/05104	A1	Pfizer Pharmaceuticals Inc.	02-04-1999	
106.	WO	99/10331	A1	Abbott Laboratories	03-04-1999	

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Sheet	7	of	12	Attorney Docket No.	PHA 4293 (3167/4Z)

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107.	WO	99/12930	A1	Glaxo Group Limited	03-18-1999
108.	WO	99/14194	A1	Merck Frosst Canada Inc.	03-25-1999
109.	WO	99/14205	A1	Almirall Prodesfarma S.A.	03-25-1999
110.	WO	99/21583	A1	Warner-Lambert Company	05-06-1999
111.	WO	99/23087	A1	Merck Frosst Canada & Co.	05-14-1999
112.	WO	99/31067	A1	Rutgers, The State University of New Jersey	06-24-1999
113.	WO	99/41241	A1	Rutgers, The State University of New Jersey	08-19-1999

#### OTHER ART - NON PATENT LITERATURE DOCUMENTS

Examiner Initials*	Cite No. <sup>1</sup>	Include name of the author (in CAPITAL LETTERS), title of the article (when appropriate), title of the item (book, magazine, journal, serial, symposium, catalog, etc.) date, page(s), volume-issue number(s), publisher, city and/or country where published.	T <sup>6</sup>
12	114.	BARNI, S., et al., Clinical Efficacy of the Aromatase Inhibitor Anastrozole in Relation to Prolactin Secretion in Heavily Pretreated Metastatic Breast Cancer, Tumori, Jan.-Feb. 1998, pp. 45-47, Vol. 84(1)	
12	115.	BENSON, J.R., et al., Tamoxifen, Biologic Therapy of Cancer: Principles and Practice, 1997, pp. 817-828, Vol. One, Ch. 29, J.F. Holland et al. (Eds.), Baltimore, MD	
12	116.	BREMBECK, F.H., et al., A Phase II Pilot Trial of 13-cis Retinoic Acid and Interferon- $\alpha$ in UICC Stage III/IV Pancreatic Cancer, Gastroenterology Oncology, April 1998, Abstract G2329, Vol. 114, No. 4, Pt. 2, A569	
12	117.	BREMBECK, F.H., et al., A Phase II Trial of 13-cis Retinoic Acid and Interferon- $\alpha$ in Patients with Advanced Pancreatic Carcinoma, Cancer, 1998, pp. 2317-2323, Vol. 83, No. 11	
12	118.	BROOKS, P.C., et al., Integrin $\alpha_v\beta_3$ Antagonists Promote Tumor Regression by Inducing Apoptosis of Angiogenic Blood Vessels, Cell, Dec. 30, 1994, pp. 1157-1164, Vol. 79	

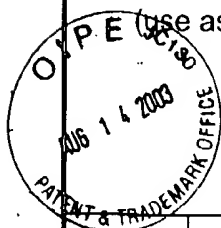
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Sheet	8	of	12	Attorney Docket No.	PHA 4293 (3167/4Z)



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<i>10</i>	119.	BROWN, F.K., et al., Matrix Metalloproteinase Inhibitors Containing a (Carboxyalkyl)amino Zinc Ligand: Modification of the P1 and P2' Residues, J. Med. Chem., 1994, pp. 674-688, Vol. 37	
<i>112</i>	120.	CAO, Y., et al., Kringle Domains of Human Angiostatin, The Journal of Biological Chemistry, Nov. 15, 1996, pp. 29461-29467, Vol. 271, No. 46	
<i>121</i>	121.	DONG, Z, et al., Macrophage-Derived Metalloelastase is Responsible for the Generation of Angiostatin in Lewis Lung Carcinoma, Cell, Mar. 21, 1997, pp. 801-810, Vol. 88	
<i>122</i>	122.	DRUMMOND, A.H., et al., Preclinical and Clinical Studies of MMP Inhibitors in Cancer, Annals of the New York Academy of Science, June 1999, pp. 228-235, Vol. 878, Robert A. Greenwald and Stanley Zucker (Eds.)	
<i>123</i>	123.	DURANDO, A., et al., Combination chemotherapy with paclitaxel (T) and epirubicin (E) for metastatic breast cancer (MBC): A phase I-II study, European Journal of Cancer, Sept. 30, 1998, pp. S12-13, Abstract 41	
<i>124</i>	124.	FAN, T.-P.D., et al., Controlling the vasculature: angiogenesis, anti-angiogenesis and vascular targeting of gene therapy, Trends Pharmacol.Sci. Feb. 1995, pp. 57-66, Vol. 16	
<i>125</i>	125.	FISHER, J.E., et al., Inhibition of Osteoclastic Bone Resorption In Vivo by Echistatin, An, "Arginyl-Glycyl-Aspartyl" (RGD)-Containing Protein, Endocrinology, 1993, pp. 1411-1413, Vol. 132, No. 3	
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Examiner Signature	<i>[Signature]</i>	Date Considered	5/18/04
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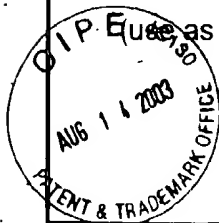
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PTO/SB/08A				<b>Complete if Known</b>	
<b>INFORMATION DISCLOSURE STATEMENT BY APPLICANT</b>  Use as many sheets as necessary)				Application Number	09/857,873
				Filing Date	October 5, 2001
				Confirmation Number	Unknown
				First Named Inventor	John P. McKearn et al.
				Group Art Unit	1614
				Examiner Name	Unknown
Sheet	9	of	12	Attorney Docket No.	PHA 4293 (3167/4Z)



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<input checked="" type="checkbox"/>	130.	HANDSCHUMACHER, R.E., et al., Purine and Pyrimidine-Antimetabolites, Chemotherapeutic Agents, pp. 712-732, Ch. XV1-2, 3rd Edition, Edited by J. Holland, et al., Lea and Febigol, publishers	
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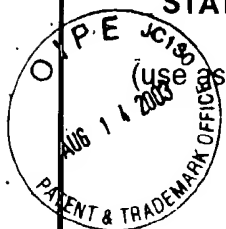
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				Group Art Unit	1614
				Examiner Name	Unknown
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# INFORMATION DISCLOSURE STATEMENT BY APPLICANT

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Application Number 09/857,873  
Filing Date October 5, 2001  
Confirmation Number Unknown  
First Named Inventor John P. McKeane et al.  
Group Art Unit 1614  
Examiner Name Unknown

Sheet 11 of 12 Attorney Docket No. PHA 4293 (3167/4Z)

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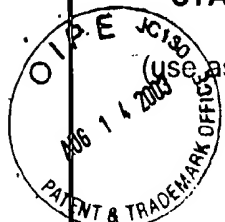
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<b>INFORMATION DISCLOSURE STATEMENT BY APPLICANT</b> (use as many sheets as necessary)			Application Number	09/857,873	
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			Confirmation Number	Unknown	
			First Named Inventor	John P. McKearn et al.	
			Group Art Unit	1614	
			Examiner Name	Unknown	
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<b>SUPPLEMENTAL          INFORMATION DISCLOSURE          STATEMENT BY APPLICANT</b>  (use as many sheets as necessary)				<b>Complete if Known</b>	
				Application Number	09/857,873
				Filing Date	October 5, 2001
				Confirmation Number	2817
				First Named Inventor	John P. McKearn et al.
				Group Art Unit	1614
Examiner Name				Rebecca Cook	
Sheet	1	of	3	Attorney Docket No. PHA 2012.1 (3167/4Z/US)	

U.S. PATENT DOCUMENTS					
Examiner Initials*	Cite No. <sup>1</sup>	U.S. Patent Document		Name of Patentee or Applicant of Cited Document	Date of Publication of Cited Document MM-DD-YYYY
		Number	Kind Code <sup>2</sup> (if known)		
<i>4</i>	172.	6,077,850		Carter et al.	06-20-2000
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FOREIGN PATENT DOCUMENTS							
Examiner Initials*	Cite No. <sup>1</sup>	Foreign Patent Document			Name of Patentee or Applicant of Cited Document	Date of Publication of Cited Document MM-DD-YYYY	T <sup>6</sup>
		Office	Number <sup>4</sup>	Kind Code <sup>2</sup> (if known)			
<i>0</i>	181.	EP	1 081 137	A1	Pfizer Products Inc.	03-07-2001	
<i>0</i>	182.	EP	1 088 550	A1	Pfizer Products Inc.	04-04-2001	

Examiner Signature	<i>W. J. [Signature]</i>	Date Considered	5/17/04
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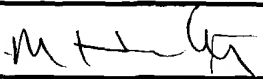
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				Examiner Name	Rebecca Cook
Sheet	2	of	3	Attorney Docket No.	PHA 2012.1 (3167/4Z/US)

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#### OTHER ART - NON PATENT LITERATURE DOCUMENTS

Examiner Initials*	Cite No. <sup>1</sup>	Include name of the author (in CAPITAL LETTERS), title of the article (when appropriate), title of the item (book, magazine, journal, serial, symposium, catalog, etc.) date, page(s), volume-issue number(s), publisher, city and/or country where published.	T <sup>6</sup>
h	196.	CRANE, C.H., M.D., et al., Initial Experience Combining Cyclooxygenase-2 Inhibition with Chemoradiation for Locally Advanced Pancreatic Cancer, Am. J. Clin. Oncol. (CCT), 2003, pp. S81-S84, Vol. 26, No. 4, Suppl. No. 2	
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Sheet	3	of	3	Attorney Docket No.	PHA 2012.1 (3167/4Z/US)

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